

International pplication No.

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|--|---|--|---------------------------------|--|
| A. CLASSIFICATION OF SUBJECT MATTER  |   |  |                                 |  |
|  | Please See Extra Sheet.<br>.514/45, 269: 586/27.21, 546/810, 562/11, 571, 561   | (/113  |                                 |  |
| According to International Patent Classification (IPC) or to both national classification and IPC  |   |  |                                 |  |
| B. FIEL  | DS SEARCHED   |  |                                 |  |
| Minimum d  | ocumentation searched (classification system followed   | I by classification symbols)   |                                 |  |
| U.S. :   | 514/45, 269; 536/27.21; 546/310; 562/11, 571; 564   | /118   |                                 |  |
| Documentat<br>searched   | ion searched other than minimum documentation to  | the extent that such documents are it  | icluded in the fields           |  |
|  | lata base consulted during the international search (r<br>Plus structure, chemical name and inventor name sea                                       |  | , search terms used)            |  |
| C. DOC   | UMENTS CONSIDERED TO BE RELEVANT  |  |                                 |  |
| Category*  | Citation of document, with indication, where ap   | propriate, of the relevant passages  | Relevant to claim No.           |  |
| Х  | WO 93/23014 A1 (Martin et al.) 25 No entire document.   | ovember 1993 (25.11.93), see   | 1-6, 20-24, 26-30, 32-34, 46-53 |  |
| X  | STOLF1 et al. Biochemical Modulat Regression of Advanced Spontaneous 5-Fluorouracil-containing Drug Combi August 1992, Vol. 52, No. 15, p document. | Murine Breast Tumors with a ination," Cancer Research, 01 ages 4074-4081, see entire | 1-6, 20-24, 26-30, 32-34, 46-53 |  |
| X Further documents or first tim the continuation of Box C. See patent family annex.   |   |  |                                 |  |
| "A" document defining the general state of the art which is not considered to be of particular series as a   |   |  |                                 |  |
| "E" earlier document positions are after the international filing date "X" document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step                           |   |  |                                 |  |
| document which mass these death in priority claim(s) or which is when the document is taken alone cited to establish the publication date of another citation or other   |   |  |                                 |  |
| document of particular relevance; the claimed invention cannot be considered to involve an inventive step when the document is combined with one or more other such documents, such combination being obvious to a person skilled in the art |   |  |                                 |  |
| "P" decument published grant to the international filing date but later "A" document member of the same patent family  |   |  |                                 |  |
| Date of the actual completion of the international search Date of mailing of the international search report   |   |  |                                 |  |
| 12 NOVEMBER 2003 25 NOV 2003   |   |  |                                 |  |
| Commissio<br>Box PCT   | Name and mailing address of the ISA IS  Commissioner of Patents and Trademarks  Box PCT  Washington, D.C. 20231  LAWRENCE ERIC CRANE                |  |                                 |  |

| C         |   |                                    |
|-----------|---|------------------------------------|
| Category* | Citation of document, with indication, where appropriate, of the relevant passages  | Relevant to claim No               |
| X         | COLOFIORE et al. Biochemical Modulation of Tumor Cell Energy IV. Evidence for the Contribution of Adenosine Triphosphate (ATP) Depletion to Chemotherapeutically Induced Tumor Regression. Biochemical Pharmacology, 27 November 1995, Vol. 50, No. 11, pages 1943-1948, see entire document. | 1-6, 20-24, 26-30,<br>32-34, 46-53 |
| X         | MARTIN et al. Marked Enhancement in vivo of Paclitaxel's (Taxol's) Tumor-Regressing Activity by ATP-Depleting Modulation. Anti-Cancer Drugs. 1996, Vol. 7, pages 655-659, see entire document.  | 1-6, 20-24, 26-30, 32-34, 46-53    |
| X         | KOUTCHER et al. Radiation Enhancement by Biochemical<br>Modulation and 5-Fluorouracil. International Journal of Radiation<br>Oncology Biology Physics. 01 December 1997, Vol. 39, No. 5, pages<br>1145-1152, see entire document.   | 1-6, 20-24, 26-30,<br>32-34, 46-53 |
| X         | GESCHWIND et al. Novel Therapy for Liver Cancer: Direct<br>Intraarterial Injection of a Potent Inhibitor of ATP Production.<br>Cancer Research. 15 July 2002, Vol. 62, pages 3909-3913, see entire<br>document.   | 1-6, 20-24, 26-30, 32-34, 46-53    |
| X         | NORI) et al. Apoptosis Induced in Advanced CD8F1-Murine<br>Mammary Tumors by the Combination of PALA, MMPR and 6AN<br>Precedes Tumor Regression and Is Preceded by ATP Depletion.<br>Cancer Chemotherapy and Pharmacology. 1997, Vol. 40, pages 376-<br>384, see entire document.             | 1-6, 20-24, 26-30,<br>32-34, 46-53 |
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Form PCT/ISA/210 (continuation of second sheet) (July 1998)\*



| A. CLASSIFICATION OF SUBJECT MATTER.  IPC (7):                                   |   |
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| A61K 31/70, 31/505; C07H 19/16; C07D 211/72, C07F 9/22; C07C 229/00; C07C 241/00 |   |
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|   | INTERNATIONAL SEARCH REPORT          | International application No. |
|---|--------------------------------------|-------------------------------|
|   | Information on patent family members | PCT/US05/16716                |
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Form PCT/ISA/210 (patent family annex) (July 1998)\*



| Box I Observations where certain claims were found unsearchable (Centinuation of item 1 of first sheet)   |
|---|
| This international report has not been established in respect of certain claims under Article 17(2)(a) for the following reasons:   |
| 1. Claims Nos.:  because they relate to subject matter not required to be searched by this Authority, namely:   |
| 2. Claims Nos.:  because they relate to parts of the international application that do not comply with the prescribed requirements to such an extent that no meaningful international search can be carried out, specifically:  |
| 3. X Claims Nos.: 7-19: 25, 31, and 35-45 because they are dependent claims and are not drafted in accordance with the second and third sentences of Rule 6.4(a).   |
| Box II Observations where unity of invention is tacking (Continuation of item 2 of first sheet)   |
| This International Searching Authority found multiple inventions in this international application, as follows:   |
|   |
| 1. As all required additional search fees were timely paid by the applicant, this international search report covers all searchable claims.  2. As all searchable claims could be searched without effort justifying an additional fee, this Authority did not invite payment |
| of any additional fee.  5. As only some of the required additional search fees were timely paid by the applicant, this international search report covers only those claims for which fees were paid, specifically claims Nos:  |
| 4. No required additional search fees were timely paid by the applicant. Consequently, this international search report is restricted to the invention first mentioned in the claims; it is covered by claims Nos.:   |
| Remark on Protest  The additional search fees were accompanied by the applicant's protest.  No protest accompanied the payment of additional search fees.   |

Form PCT/ISA/210 (continuation of first sheet(1)) (July 1998)\*

| Box III TEXT OF THE ABSTRACT (Sectionation of item 5 of the first elect) |           | INTERNATIONAL        | <b>LEARCH</b> | REPORT                           | International application No. |
|--|-----------|----------------------|---------------|----------------------------------|-------------------------------|
|  |           |                      |               |                                  | PCT/US03/18716                |
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|  | Box III 7 | TEXT OF THE ABSTRACT | (Canuatio     | on of item 5 of the first sheet) |                               |
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